Figure 1

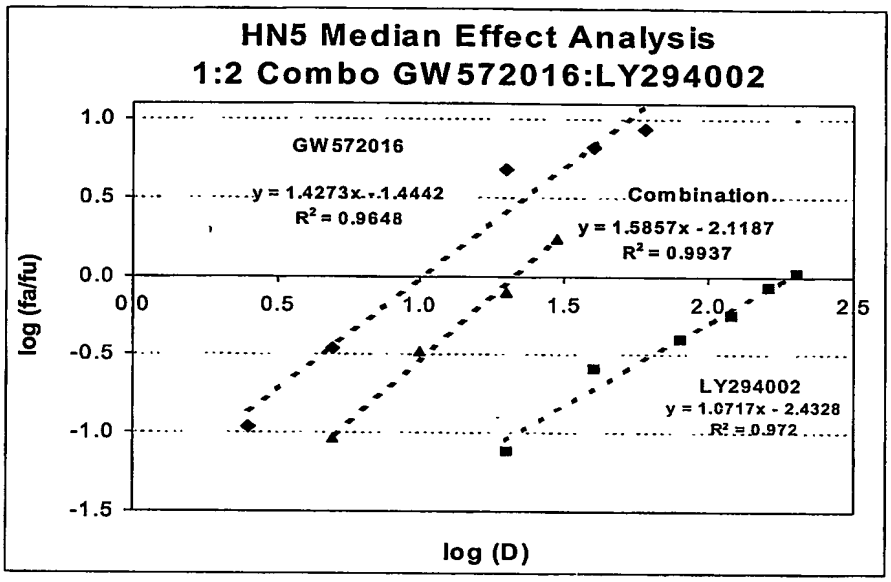

HN5 Median Effect Analysis
1:2 Combo GW572016:LY294002
GW572016
y = 1.4273x - 1.4442
R² = 0.9648
Combination
y = 1.5857x - 2.1187
R² = 0.9937
LY294002
y = 1.0717x - 2.4328
R² = 0.972
log (fa/fu)
log (D)
1.0
0.5
0.0
-0.5
-1.0
-1.5
0.0
0.5
1.0
1.5
2.0
2.5

Figure 2

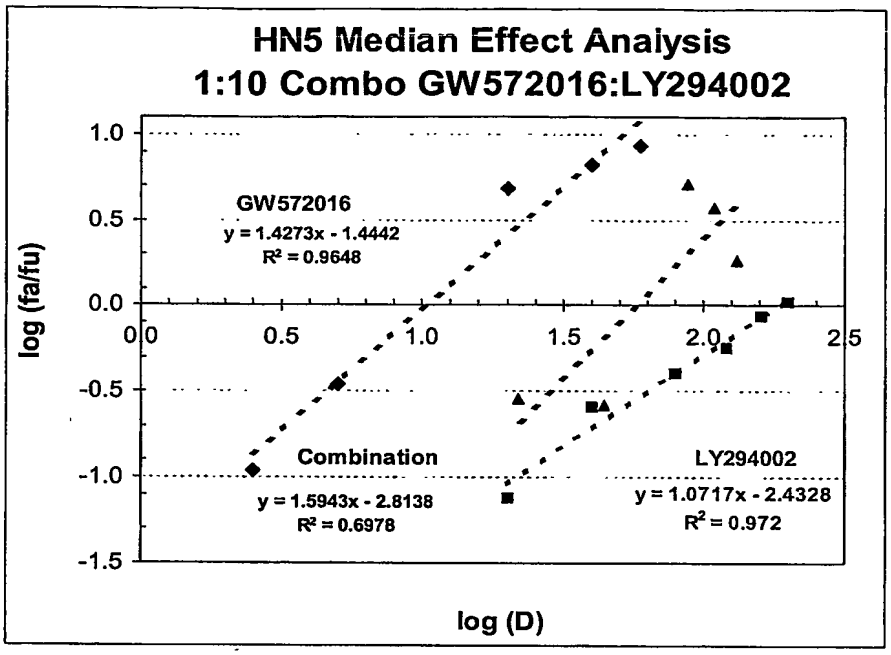

HN5 Median Effect Analysis
1:10 Combo GW572016:LY294002
log (fa/fu)
log (D)
1.0
0.5
0.0
-0.5
-1.0
-1.5
0.0
0.5
1.0
1.5
2.0
2.5
GW572016
y = 1.4273x - 1.4442
R² = 0.9648
Combination
y = 1.5943x - 2.8138
R² = 0.6978
LY294002
y = 1.0717x - 2.4328
R² = 0.972

Figure 3

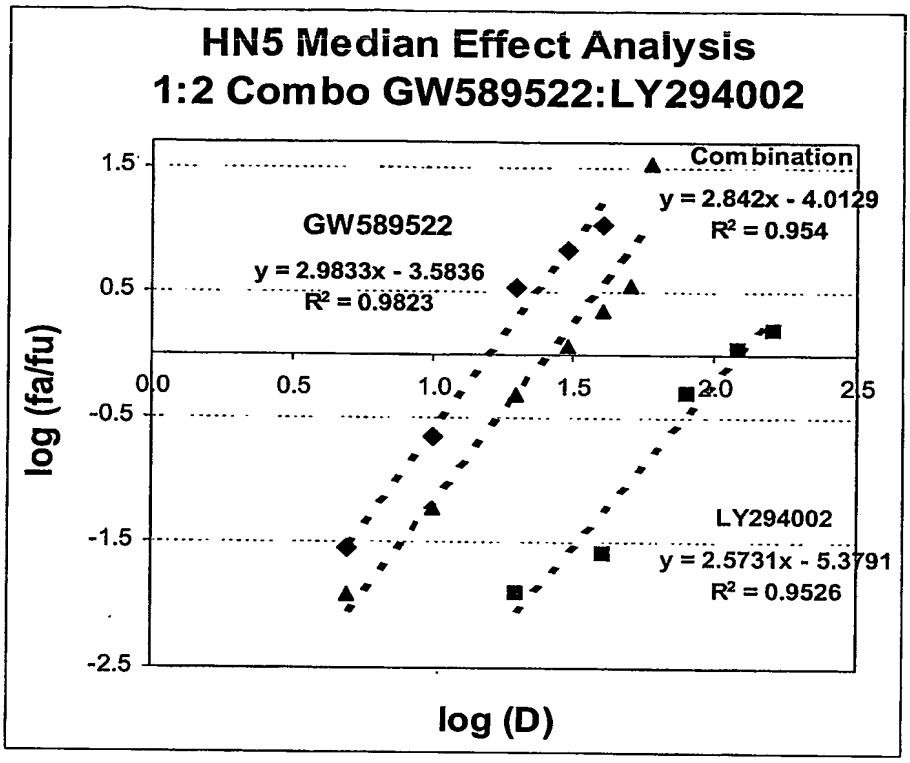

HN5 Median Effect Analysis
1:2 Combo GW589522:LY294002
Combination
y = 2.842x - 4.0129
R² = 0.954
GW589522
y = 2.9833x - 3.5836
R² = 0.9823
LY294002
y = 2.5731x - 5.3791
R² = 0.9526
log (fa/fu)
log (D)
1.5
0.5
-0.5
-1.5
-2.5
0.0
0.5
1.0
1.5
2.0
2.5

Figure 4

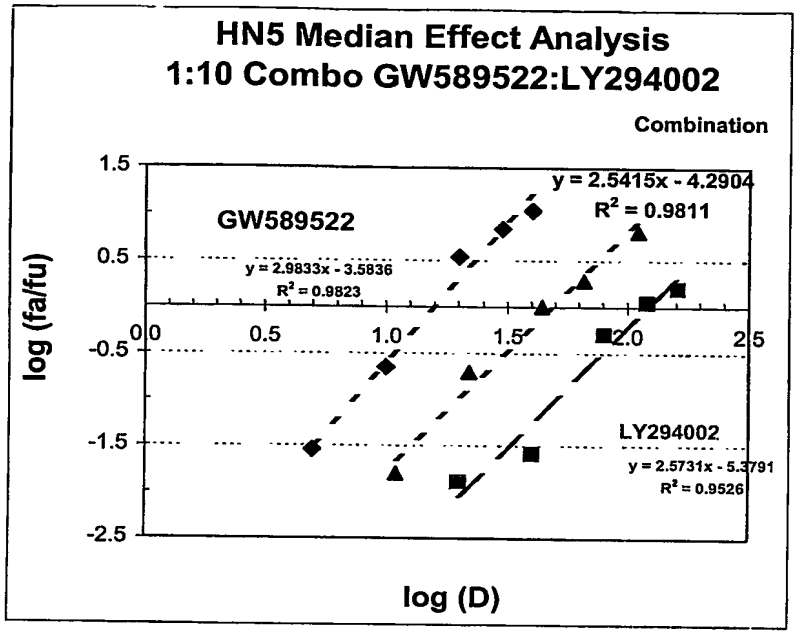

HN5 Median Effect Analysis
1:10 Combo GW589522:LY294002
Combination
GW589522
y = 2.9833x - 3.5836
R² = 0.9823
y = 2.5415x - 4.2904
R² = 0.9811
LY294002
y = 2.5731x - 5.3791
R² = 0.9526
log (fa/fu)
log (D)
1.5
0.5
-0.5
-1.5
-2.5
0.0
0.5
1.0
1.5
2.0
2.5

Figure 5

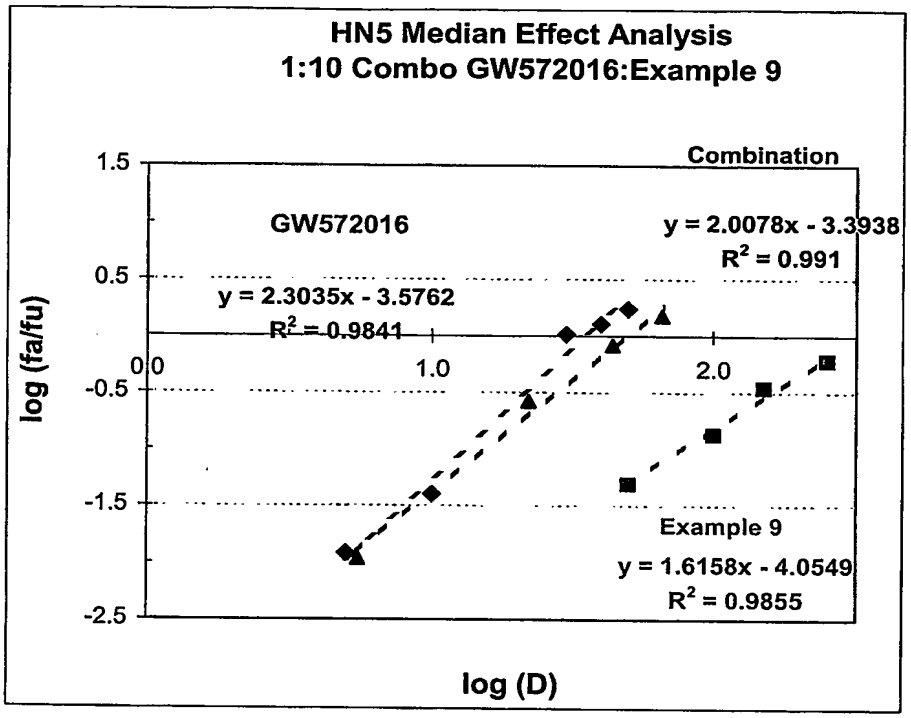

HN5 Median Effect Analysis
1:10 Combo GW572016:Example 9
Combination
GW572016
y = 2.0078x - 3.3938
R2 = 0.991
y = 2.3035x - 3.5762
R2 = 0.9841
Example 9
y = 1.6158x - 4.0549
R2 = 0.9855
log (fa/fu)
log (D)
1.5
0.5
-0.5
-1.5
-2.5
0.0
1.0
2.0

Figure 6

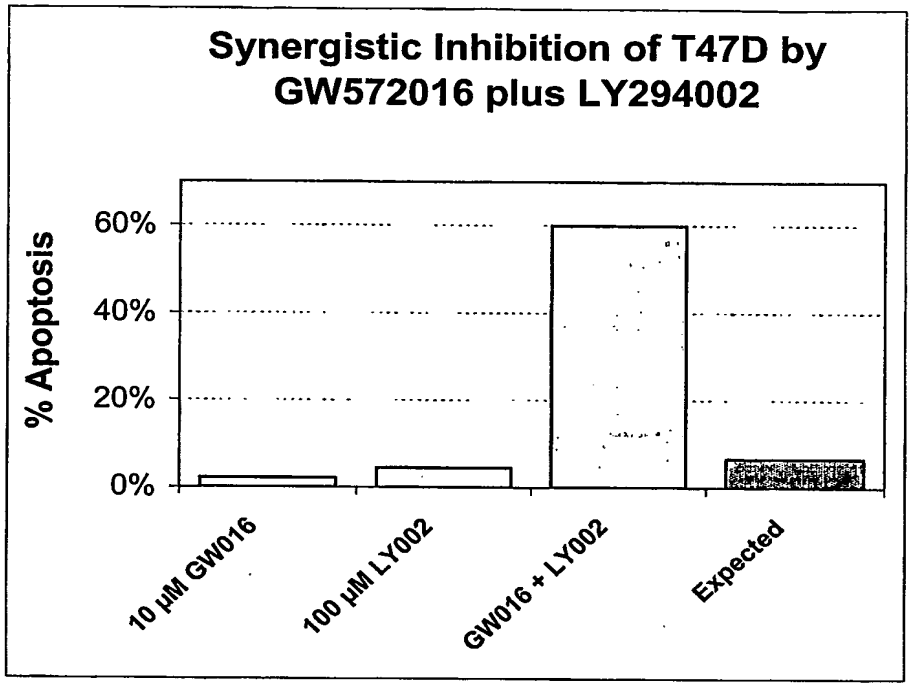
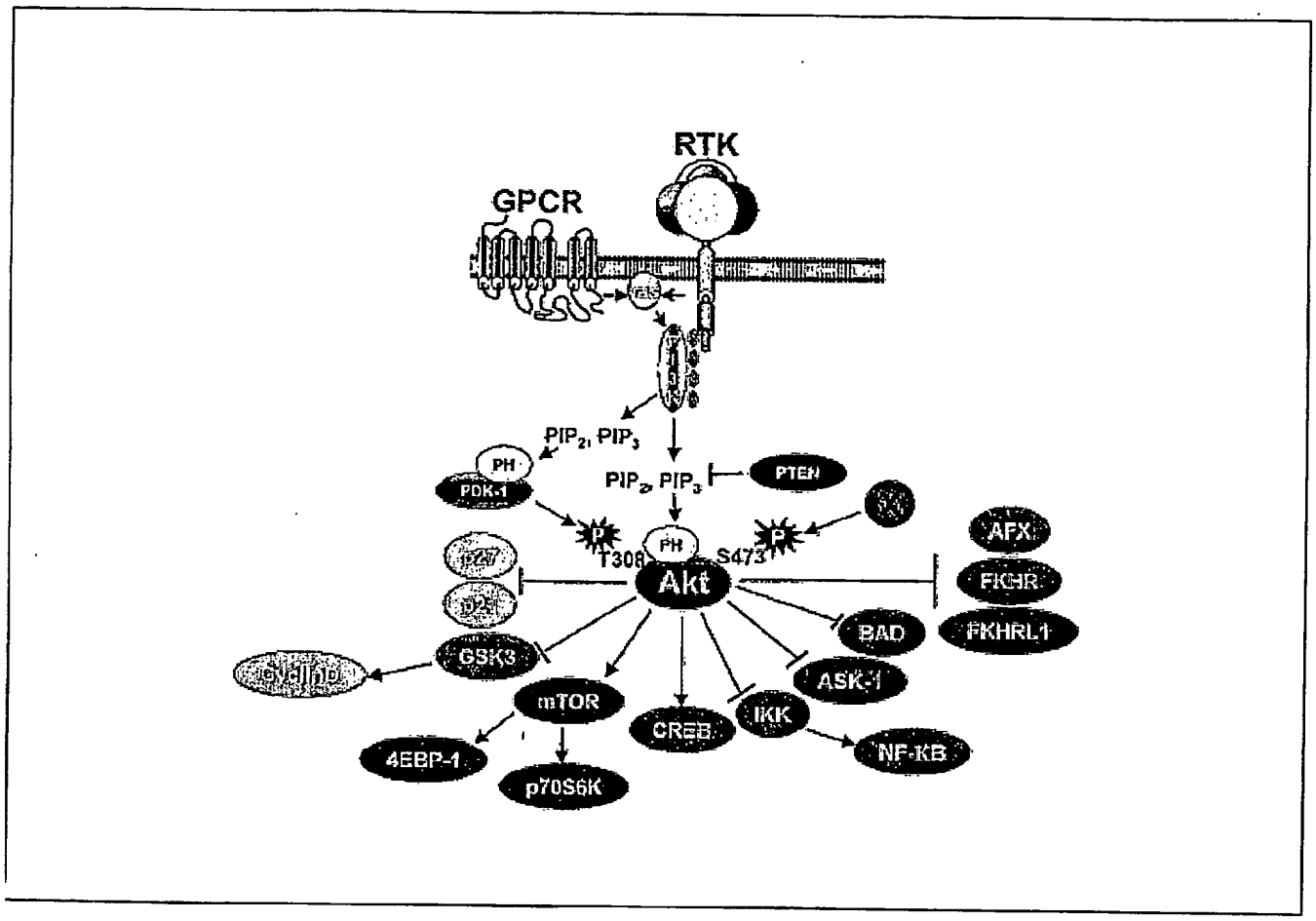

Synergistic Inhibition of T47D by GW572016 plus LY294002
% Apoptosis
60%
40%
20%
0%
10 µM GW016
100 µM LY002
GW016 + LY002
Expected

Figure 7


RTK
GPCR
PIP2, PIP3
PH
PDK-1
PIP2, PIP3
PTEN
P
T308
PH
S473
P
AFX
FKHR
FKHRL1
p27
p21
Akt
BAD
GSK3
Cyclin D
ASK-1
mTOR
CREB
IKK
NF-KB
4EBP-1
p70S6K